PCT/US2004/011168

12 April 2004

10 April 2003

## **Amendments to the Claims**

This listing of claims will replace all prior versions, and listings, of claims in the application:

## **Listing of Claims:**

- 1. (ORIGINAL) A device comprising an extracellular matrix having an internal pH between 4.0 and 6.0, wherein the extracellular matrix contains heparin or a heparin-related compound bound to a protein, wherein the protein has a pH dependent binding to the heparin or the heparin-related compound.
- 2. (ORIGINAL) The device of claim 1, wherein the internal pH is between 5 and 6.
  - 3. (ORIGINAL) The device of claim 1, wherein the internal pH is about 5.5.
  - 4. (ORIGINAL) The device of claim 1, wherein the bound protein is VEGF.
- 5. (CURRENTLY AMENDED) The device of <u>claim 1 elaims 1 or 4</u>, wherein the extracellular matrix further comprises fibronectin or a fibronectin fragment that binds to the bound protein.
  - 6. (CANCEL)
- 7. (ORIGINAL) The device of claim 1, wherein the heparin-related compound is heparan sulfate or heparan sulfate proteoglycan.
- 8. (CURRENTLY AMENDED) The device of <u>claim 1-elaims 1, 4, 5, 6 or 7</u>, wherein the bound protein contains a heparin-binding consensus sequence.
- 9. (ORIGINAL) The device of claim 8, wherein the heparin-binding consensus sequence is XBBBXXBX or XBBXBX, where B is a basic amino acid residue or His and X is any amino acid residue.
- 10. (CURRENTLY AMENDED) The device of claim 1 claims 1, 4, 5, 6 or 7, wherein the bound protein contains a glycine-like box, wherein said glycine-like box is from about seven to twelve amino acids and contains at least two Gly residues and two-five basic amino acid residues.

INTERNATIONAL APPL. NO.

12 April 2004

10 April 2003

- 11. (ORIGINAL) The device of claim 1, wherein the glycine-like box is SEO ID NO:1.
- 12. (ORIGINAL) The device of claim 11, wherein the bound protein is VEGF<sub>121</sub> or VEGF<sub>165</sub>.
- 13. (CURRENTLY AMENDED) The device of claim1 claims 1, 4, 5, 6, 7, 8, 9, 1-. 11 or 12, wherein the device is formed in situ in a subject.
- 14. (CURRENTLY AMENDED) A kit for making the device of claim 1 elaims 1, 4, 5, 6, 7, 8, 9, 10, 11, 12, or 13, wherein the kit contains a vial containing heparin or a heparinrelated compound and a second vial containing fibronectin or a protein containing a heparinbind domain.
- 15. (CURRENTLY AMENDED) The device of claim 1 claims 1, 4, 5, 6, 7, 8, 9, 10, 11, or 12, wherein the extracellular matrix is attached to or encased within a compound selected from the group consisting of a film, a hydrocolloid, a hydrogel, a foam, a gelatin, a bead, a bandage, and a cellophane.
- 16. (CURRENTLY AMENDED) The device method of claim 1-elaim 6, wherein the heparin-related compound is a heparin-related oligiosaccharide of 8-16 sugars.
- 17. (CURRENTLY AMENDED) A method of stimulating angiogenesis at a clinically relevant site in a mammal, said method comprising administering an effective amount of the device of claim 1 elaims 1, 4, 5, 6, 7, 8, 9, 10, 11, 12, 13 or 15 capable of releasing a pharmaceutically effective amount of the bound protein at said clinically relevant site.
- (CURRENTLY AMENDED) The method of claim 17 claims 5 and 17, wherein 18. said clinically relevant site is a wound and the bound protein is an angiogenic protein that promotes wound healing.
- 19. (CURRENTLY AMENDED) The method of claim 17-claims 17 or 18, where the bound protein is VEGF.
- 20. (ORIGINAL) The method of claim 19, wherein the VEGF is VEGF<sub>121</sub> or VEGF<sub>165</sub>.

PCT/US2004/011168

10 April 2003

- .21. (CURRENTLY AMENDED) The method of claim17-claims 17, 18, 19 or 20, wherein the device is administered by injection or surgical placement.
- 22. (CURRENTLY AMENDED) The method of claim 17 elaims 17, 18, 19, 20 or 21, wherein the device is formed in the mammal in situ.
- 23. (NEW) The device of claim 4, wherein the extracellular matrix further comprises fibronectin or a fibronectin fragment that binds to the bound protein.
- (NEW) The device of 4, wherein the bound protein contains a heparin-24. binding consensus sequence.
- 25. (NEW) The device of 4, wherein the bound protein contains a glycine-like box, wherein said glycine-like box is from about seven to twelve amino acids and contains at least two Gly residues and two-five basic amino acid residues.
- 26. (NEW) The device of 7, wherein the bound protein contains a glycine-like box, wherein said glycine-like box is from about seven to twelve amino acids and contains at least two Gly residues and two-five basic amino acid residues.
  - 27. (NEW) The device of 4, wherein the device is formed in situ in a subject.
  - 28. (NEW) The device of 7, wherein the device is formed in situ in a subject.
  - (NEW) The device of 12, wherein the device is formed in situ in a subject. 29.
- 30. (NEW) The method of claim 19, wherein the extracellular matrix further comprises fibronectin or a fibronectin fragment that binds to the bound protein and wherein the heparin-related compound is heparan sulfate or heparan sulfate proteoglycan.